# **Hit List**

Clear Generate Collection Print Fwd Refs Bkwd Refs
Generate OACS

## Search Results - Record(s) 1 through 6 of 6 returned.

☐ 1. Document ID: US 20030150008 A1

Using default format because multiple data bases are involved.

L3: Entry 1 of 6

File: PGPB

Aug 7, 2003

PGPUB-DOCUMENT-NUMBER: 20030150008

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030150008 A1

TITLE: Transgenic plants containing altered levels of steroid compounds

PUBLICATION-DATE: August 7, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Karunanandaa, Balasulojini	St. Louis	MO	US	
Post-Beittenmiller, Martha	St. Louis	MO	US	
Venkatramesh, Mylavarapu	St. Louis	MO	US	
Kishore, Ganesh M.	St. Louis	MO	US	
Thorne, Gregory M.	St. Louis	MO	US	
LeDeaux, John R.	St. Louis	MO	US	

US-CL-CURRENT: 800/278; 435/189, 435/410, 536/23.2

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWAC	Drawt De
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L3: I	Entry	2 of 6				]	File: P	GPB		Jul	3,	2003

PGPUB-DOCUMENT-NUMBER: 20030125573

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030125573 A1

TITLE: Method of vitamin production

PUBLICATION-DATE: July 3, 2003

INVENTOR-INFORMATION:

NAME CITY STATE COUNTRY RULE-47

Millis, James R. Kohler WI US

h e b b g e e e f b e

Saucy, Gabriel G.
Maurina-Brunker, Julie
McMullin, Thomas W.

Essex Fells NY US
Appleton WI US
Manitowoc WI US

US-CL-CURRENT: 549/411; 435/155

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draw. De

☐ 3. Document ID: US 20030092144 A1

L3: Entry 3 of 6

File: PGPB

May 15, 2003

RULE-47

PGPUB-DOCUMENT-NUMBER: 20030092144

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030092144 A1

TITLE: Production of farnesol and geranylgeraniol

PUBLICATION-DATE: May 15, 2003

INVENTOR-INFORMATION:

COUNTRY STATE CITY NAME US WI Millis, James R. Kohler US Appleton WI Maurina-Brunker, Julie ÚS WΙ Manitowoc McMullin, Thomas W.

US-CL-CURRENT: 435/157; 435/252.3

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draw. De

4. Document ID: US 20020108148 A1

L3: Entry 4 of 6

File: PGPB

- Aug 8, 2002

PGPUB-DOCUMENT-NUMBER: 20020108148

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020108148 A1

TITLE: Nucleic acid sequences to proteins involved in isoprenoid synthesis

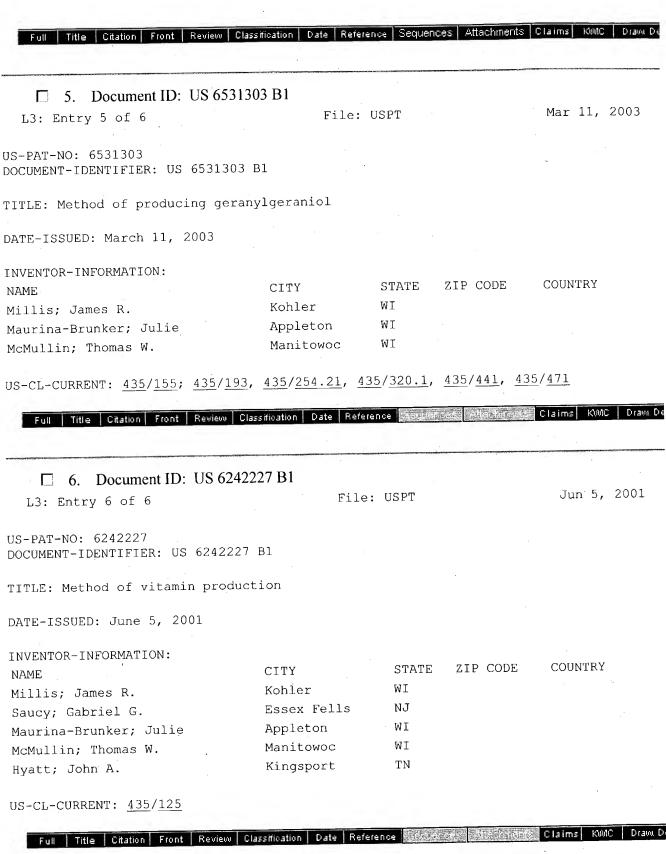
PUBLICATION-DATE: August 8, 2002

INVENTOR-INFORMATION:

NAME CITY STATE COUNTRY RULE-47
Boronat, Albert Barcelona MO ES
Campos, Narciso Barcelona ES

Kishore, Ganesh M. Creve Coeur US

US-CL-CURRENT: 800/284; 435/189, 435/320.1, 435/410, 435/69.1, 536/23.2



. 1	Full	Title	Citation	Front	Review	Classification	Date	Reference	2012/01/2012/2012		Flams	Konc	DIAMA De
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Terms	Documents
1-deoxy-D-xylulose 5 phosphate reductoisomerase.clm.	6

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Previous Page Next Page Go to Doc#

## First Hit



L3: Entry 4 of 6

File: PGPB

Aug 8, 2002

PGPUB-DOCUMENT-NUMBER: 20020108148

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020108148 A1

TITLE: Nucleic acid sequences to proteins involved in isoprenoid synthesis

PUBLICATION-DATE: August 8, 2002

#### INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Boronat, Albert	Barcelona	MO	ES	
Campos, Narciso	Barcelona		ES	
Kishore, Ganesh M.	Creve Coeur		US	

US-CL-CURRENT: 800/284; 435/189, 435/320.1, 435/410, 435/69.1, 536/23.2

#### CLAIMS:

What is claimed is:

- 1. An isolated nucleic acid sequence encoding  $\frac{1-\text{deoxy-D-xylulose }5-\text{phosphate}}{\text{reductoisomerase}}$  from a eukaryotic source.
- 2. An isolated nucleic acid sequence of claim 1, wherein said nucleic acid sequence is isolated from a plant source.
- 3. An isolated nucleic acid sequence of claim 2, wherein said nucleic acid sequence is isolated from Arabidopsis.
- 4. An isolated polynucleotide selected from the group consisting of: a) an isolated polynucleotide comprising a nucleotide sequence encoding the polypeptide of SEQ ID NO:2; b) an isolated polynucleotide comprising SEQ ID NO:1; c) an isolated polynucleotide comprising SEQ ID NO:1; c) an isolated polynucleotide comprising a nucleotide sequence which has at least 70% identity to that of SEQ ID NO:1 over the entire length of SEQ ID NO:1; d) an isolated polynucleotide comprising a nucleotide sequence which has at least 80% identity to that of SEQ ID NO:1 over the entire length of SEQ ID NO:1; e) an isolated polynucleotide comprising a nucleotide sequence which has at least 90% identity to that of SEQ ID NO:1 over the entire length of SEQ ID NO:1; f) an isolated polynucleotide comprising a nucleotide sequence which has at least 95% identity to that of SEQ ID NO:1 over the entire length of SEQ ID NO:1; g) an isolated polynucleotide that hybridizes, under stringent conditions, to SEQ ID NO:1 or a fragment thereof; and h) an isolated polynucleotide complementary to the polynucleotide sequence of (a), (b), (c), (d), (e), (f), or (g).
- 5. A DNA construct, comprising; as operably associated components in the 5' to 3' direction of transcription, a promoter functional in a plant cell, a nucleic acid sequence encoding <a href="1-deoxy-D-xylulose-5-phosphate reductoisomerase">1-deoxy-D-xylulose-5-phosphate reductoisomerase</a>, and a transcriptional termination sequence.

- 6. The DNA construct according to claim 5, wherein said nucleic acid sequence is isolated from a eukaryotic source.
- 7. The DNA construct according to claim 5, wherein said nucleic acid sequence is isolated from a plant source.
- 8. The DNA construct according to claim 5, wherein said nucleic acid sequence is isolated from Arabidopsis.
- 9. A host cell comprising the construct of claim 5.
- 10. A host cell according to claim 9, wherein the host cell is a plant cell.
- 11. A plant comprising a cell according to claim 10.
- 12. A method for the alteration of the isoprenoid content in a plant, comprising; transforming said host plant with a construct comprising as operably linked components, a transcriptional initiation region functional in a plant, a nucleic acid sequence encoding 1-deoxy-D-xylulose 5-phosphate reductoisomerase, and a transcriptional termination region.
- 13. A method for the alteration of the isoprenoid content in a plant according to claim 12, wherein said nucleic acid sequence is in the sense orientation.
- 14. A method according to claim 13, wherein the isoprenoid content is increased.
- 15. A method for the alteration of the isoprenoid content in a plant according to claim 12, wherein said nucleic acid sequence is in the antisense orientation.
- 16. A method according to claim 15, wherein the isoprenoid content is decreased.
- 17. A method for producing an isoprenoid compound of interest in a plant cell, said method comprising obtaining a transformed plant, said plant having and expressing in its genome: a primary construct comprising a DNA sequence encoding 1-deoxy-D-xylulose 5-phosphate reductoisomerase operably linked to a transcriptional initiation region functional in a plant cell; and, at least one secondary construct comprising a DNA sequence encoding a protein involved in the production of a particular isoprenoid operably linked to a transcriptional initiation region functional in a plant cell.
- 18. A method according to claim 17, wherein said protein is involved in the production of isoprenoids selected from the group consisting of tocopherols, carotenoids, monoterpenes, diterpenes, and plastoquinones.
- 19. A method for increasing the non-mevalonate isoprenoid biosynthetic flux in cell from a host plant, said method comprising transforming said host plant with a construct comprising as operably linked components, a transcriptional initiation region functional in a plant, a DNA coding <a href="1-deoxy-D-xylulose-5-phosphate">1-deoxy-D-xylulose-5-phosphate</a> reductoisomerase, and a transcriptional termination region.
- 20. A method for modulating disease resistance in a plant, comprising: growing a plant which contains in its genome a construct which provides for expression of a 1-deoxy-D-xylulose 5-phosphate reductoisomerase gene.

# **Hit List**

Clear	Generate Collection	Print	Fwd Refs	Bkwd Refs
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# Search Results - Record(s) 11 through 19 of 19 returned.

☐ 11. Document ID: US 20020108148 A1

L2: Entry 11 of 19

File: PGPB

Aug 8, 2002

PGPUB-DOCUMENT-NUMBER: 20020108148

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020108148 A1

TITLE: Nucleic acid sequences to proteins involved in isoprenoid synthesis

PUBLICATION-DATE: August 8, 2002

INVENTOR-INFORMATION:

NAME CITY STATE COUNTRY RULE-47

Boronat, Albert Barcelona MO ES
Campos, Narciso Barcelona ES
Kishore, Ganesh M. Creve Coeur US

US-CL-CURRENT: 800/284; 435/189, 435/320.1, 435/410, 435/69.1, 536/23.2

Ī	Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw, De
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### ☐ 12. Document ID: US 20020069426 A1

L2: Entry 12 of 19

File: PGPB

Jun 6, 2002

PGPUB-DOCUMENT-NUMBER: 20020069426

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020069426 A1

TITLE: Methyl-D-erythritol phosphate pathway genes

PUBLICATION-DATE: June 6, 2002

INVENTOR-INFORMATION:

STATE COUNTRY RULE-47 CITY NAME Barcelona MO ES Boronat, Albert Barcelona MO ES Campos, Narciso ES MO Barcelona Rodriguez-Concepcion, Manual FR Rohmer, Michel Strasbourg FRRixheim Seeman, Myriam

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Record List Display

Valentin, Henry E.

Chesterfield

US

Venkatesh, Tyamagondlu V.

St. Louis

US

Venkatramesh, Mylavarapu

Ballwin

US

US-CL-CURRENT: 800/278; 435/69.8, 530/370, 536/23.6, 800/286

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draws

☐ 13. Document ID: US 6531303 B1

L2: Entry 13 of 19

File: USPT

Mar 11, 2003

US-PAT-NO: 6531303

DOCUMENT-IDENTIFIER: US 6531303 B1

TITLE: Method of producing geranylgeraniol

DATE-ISSUED: March 11, 2003

INVENTOR-INFORMATION:

NAME

CITY

ZIP CODE STATE

COUNTRY

Millis; James R.

Kohler

WI

Maurina-Brunker; Julie

WΙ Appleton

McMullin; Thomas W.

Manitowoc

US-CL-CURRENT: 435/155; 435/193, 435/254.21, 435/320.1, 435/441, 435/471

Full Title Citation Front Review Classification Date Reference Constitution Ett. March Claims KWIC Draw. De

☐ 14. Document ID: US 6420159 B2

L2: Entry 14 of 19

File: USPT

Jul 16, 2002

US-PAT-NO: 6420159

DOCUMENT-IDENTIFIER: US 6420159 B2

TITLE: 1-deoxy-D-xylulose-5-phosphate reductoisomerases, and methods of use

DATE-ISSUED: July 16, 2002

INVENTOR-INFORMATION:

NAME

CITY

STATE

ZIP CODE

COUNTRY

Croteau; Rodney B.

Pullman

WΑ

Lange; Bernd M.

Pullman

WA

US-CL-CURRENT: 435/233

Full Title Citation Front Review Classification Date Reference Reference Citation Claims KMC Draw De

cc

☐ 15. Document ID: US 6387637 B1

L2: Entry 15 of 19

File: USPT

STATE

ZIP CODE

May 14, 2002

COUNTRY

US-PAT-NO: 6387637

DOCUMENT-IDENTIFIER: US 6387637 B1

TITLE: Herbicide target genes and method

DATE-ISSUED: May 14, 2002

INVENTOR-INFORMATION:

NAME CITY

Levin; Joshua Z. Durham NC Budziszewski; Gregory J. Durham NC

Potter; Sharon L. Raleigh NC Wegrich; Lynette M. San Jose CA

US-CL-CURRENT: 435/7.1; 530/350, 530/380

Full Title Citation Front Review Classification Date Reference Section Section (Claims KWC Draw De

☐ 16. Document ID: US 6281017 B1

L2: Entry 16 of 19

File: USPT

Aug 28, 2001

US-PAT-NO: 6281017

DOCUMENT-IDENTIFIER: US 6281017 B1

TITLE: 1-deoxy-d-xylulose-5-phosphate reductoisomerases and method of use

DATE-ISSUED: August 28, 2001

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Croteau; Rodney B. Pullman WA Lange; Bernd M. Pullman WA

US-CL-CURRENT:  $\underline{435/468}$ ;  $\underline{435/189}$ ,  $\underline{435/233}$ ,  $\underline{435/320.1}$ ,  $\underline{435/410}$ ,  $\underline{435/476}$ 

☐ 17. Document ID: US 6242227 B1

L2: Entry 17 of 19

File: USPT

Jun 5, 2001

US-PAT-NO: 6242227

DOCUMENT-IDENTIFIER: US 6242227 B1

Jun 15, 2000

Record List Display

TITLE: Method of vitamin production

DATE-ISSUED: June 5, 2001

INVENTOR-INFORMATION:

ZIP CODE COUNTRY STATE CITY NAME WI Kohler Millis; James R. Essex Fells N.T Saucy; Gabriel G. WI Appleton Maurina-Brunker; Julie WI Manitowoc McMullin; Thomas W. Kingsport TNHyatt; John A.

US-CL-CURRENT: 435/125

Full Title	Citation	Front	Review	Classification	Date	Reference	े के <b>अगर्क</b> शिक्षणिक स्ट्र	Claims	KOMC	Draw, De
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□ 18.	Docum	ent II	): US 2	2002010814	18 A I					
L2: Entry	7 18 of	19		-		File:	DWPI	Aug	8,	2002

DERWENT-ACC-NO: 2003-066660

DERWENT-WEEK: 200306

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TITLE: New nucleic acid sequence encoding 1-deoxy-D-xylulose 5-phosphate

reductoisomerase from an eukaryotic source, useful for altering isoprenoid content

and composition, and modulating disease resistance in plants

INVENTOR: BORONAT, A; CAMPOS, N; KISHORE, G M

PRIORITY-DATA: 2001US-0987025 (November 13, 2001), 1999US-129899P (April 15, 1999),

1999US-146461P (July 30, 1999), 2000US-0549787 (April 14, 2000)

PATENT-FAMILY:

 PUB-NO
 PUB-DATE
 LANGUAGE
 PAGES
 MAIN-IPC

 US 20020108148 A1
 August 8, 2002
 019
 A01H005/00

INT-CL (IPC): A01  $\underline{H}$  5/00; C07  $\underline{H}$  21/04; C12  $\underline{N}$  5/04; C12  $\underline{N}$  9/02; C12  $\underline{P}$  21/02

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File: DWPI

DERWENT-ACC-NO: 2000-431295

DERWENT-WEEK: 200037

L2: Entry 19 of 19

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TITLE: Novel polynucleotide encoding isopentenyl diphosphate biosynthetic enzymes useful for producing transgenic plants with altered isopentenyl diphosphate levels

and for selecting polynucleotides affecting expression of the enzyme

INVENTOR: CAHOON, R E; LEE, J ; TAO, Y

PRIORITY-DATA: 1998US-110865P (December 4, 1998)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
WO 200034448 A1	June 15, 2000	. E	063	C12N009/04
AU 200021633 A	June 26, 2000		000	C12N009/04
EP 1135471 A1	September 26, 2001	Ε	000	C12N009/04

INT-CL (IPC):  $\underline{\text{C12}}\ \underline{\text{N}}\ \underline{9/04};\ \underline{\text{C12}}\ \underline{\text{N}}\ \underline{15/63};\ \underline{\text{C12}}\ \underline{\text{N}}\ \underline{15/67};\ \underline{\text{C12}}\ \underline{\text{N}}\ \underline{15/82};\ \underline{\text{C12}}\ \underline{\text{N}}\ \underline{15/86}$ 

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Previous Page Next Page Go to Doc#

## First Hit Fwd Refs

Generate Collection Print

L2: Entry 16 of 19

File: USPT

Aug 28, 2001

US-PAT-NO: 6281017

DOCUMENT-IDENTIFIER: US 6281017 B1

TITLE: 1-deoxy-d-xylulose-5-phosphate reductoisomerases and method of use

DATE-ISSUED: August 28, 2001

INVENTOR-INFORMATION:

NAME

CITY

STATE

ZIP CODE

COUNTRY

Croteau; Rodney B.

Pullman

WA WA

Lange; Bernd M.

Pullman

US-CL-CURRENT: 435/468; 435/189, 435/233, 435/320.1, 435/410, 435/476

#### CLAIMS:

#### What is claimed is:

- 1. An isolated nucleic acid molecule that hybridizes under stringent conditions to the nucleic acid molecule of SEQ ID NO:1, or to the complement of the nucleic acid molecule of SEQ ID NO:1, provided that said isolated nucleic acid molecule does not consist of a nucleic acid sequence selected from the group consisting of SEQ ID NO:10 and SEQ ID NO:11 or a nucleic acid sequence complementary to a nucleic acid sequence selected from the group consisting of SEQ ID NO:10 and SEQ ID NO:11, wherein said stringent hybridization conditions consist of hybridization in 3.times.SSC at 65.degree. C. for 16 hours, followed by two washes in 2.times.SSC at 23.degree. C. for 20 minutes per wash, followed by one wash in 0.5.times.SSC at 55.degree. C. for 30 minutes.
- 2. An isolated nucleic acid molecule of claim 1 wherein said isolated nucleic acid molecule encodes a plant 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein.
- 3. An isolated nucleic acid molecule of claim 1 wherein said isolated nucleic acid molecule encodes an essential oil plant 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein.
- 4. An isolated nucleic acid molecule of claim 3 wherein said isolated nucleic acid molecule encodes a Mentha 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein.
- 5. An isolated nucleic acid molecule of claim 1 wherein said nucleic acid molecule encodes a 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein comprising the amino acid sequence set forth in SEQ ID NO:2.
- 6. An isolated nucleic acid molecule of claim 1 comprising the nucleic acid sequence of SEQ ID NO:1.

- 7. A replicable vector comprising a first nucleic acid molecule that hybridizes under stringent conditions to a second nucleic acid molecule consisting of the nucleic acid sequence set forth in SEQ ID NO:1, or to a third nucleic acid molecule consisting of the complement of the nucleic acid sequence set forth in SEQ ID NO:1, provided that said first nucleic acid molecule does not consist of a nucleic acid sequence selected from the group consisting of SEQ ID NO:10 and SEQ ID NO:11 or a nucleic acid sequence complementary to a nucleic acid sequence selected from the group consisting of SEQ ID NO:10 and SEQ ID NO:11, wherein said stringent hybridization conditions consist of hybridization in 3.times.SSC at 65.degree. C. for 16 hours, followed by two washes in 2.times.SSC at 23.degree. C. for 20 minutes per wash, followed by one wash in 0.5.times.SSC at 55.degree. C. for 30 minutes.
- 8. A replicable vector of claim 7 wherein said first nucleic acid molecule encodes a plant 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein.
- 9. A replicable vector of claim 7 wherein said first nucleic acid molecule encodes a Mentha 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein.
- 10. A replicable vector of claim 7 wherein said first nucleic acid molecule encodes a 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein comprising the amino acid sequence set forth in SEQ ID NO:2.
- 11. A replicable vector of claim 7 wherein said first nucleic acid molecule comprises the nucleic acid sequence set forth in SEQ ID NO:1, or the complement of the nucleic acid sequence set forth in SEQ ID NO:1.
- 12. A host cell comprising a vector of claim 7.
- 13. A host cell comprising a vector of claim 11.
- 14. A host cell of claim 12 wherein said host cell is a plant cell.
- 15. A host cell of claim 13 wherein said host cell is a plant cell.
- 16. A method of enhancing the level of expression of 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein in a host cell comprising introducing into said host cell a replicable expression vector comprising a nucleic acid molecule that encodes a 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein under conditions that enable expression of said protein in said host cell, wherein said nucleic acid molecule hybridizes to the complement of a nucleic acid molecule consisting of the nucleic acid sequence set forth in SEQ ID NO:1 under hybridization conditions consisting of hybridization in 3.times.SSC at 65.degree. C. for 16 hours, followed by two washes in 2.times.SSC at 23.degree. C. for 20 minutes per wash, followed by one wash in 0.5.times.SSC at 55.degree. C. for 30 minutes.
- 17. A method of reducing the level of expression of 1-deoxy-D-xylulose-5-phosphate reductoisomerase protein in a host cell comprising introducing into said host cell a replicable expression vector comprising a nucleic acid molecule that expresses an RNA molecule that hybridizes under stringent conditions to the nucleic acid sequence of SEQ ID NO:1, wherein said stringent hybridization conditions consist of hybridization in 3.times.SSC at 65.degree. C. for 16 hours, followed by two washes in 2.times.SSC at 23.degree. C. for 20 minutes per wash, followed

h

by one wash in 0.5.times.SSC at 55.degree. C. for 30 minutes.

# **WEST Search History**

Hide Items Restore Clear Cancel

DATE: Tuesday, January 13, 2004

Hide?	Set Nam	e Query	Hit Count
	DB=PG	SPB, USPT, USOC, EPAB, JPAB, DWPI; PLUR = YES; OP = ACCEPTANCE OF STATES O	ADJ
	L3	1-deoxy-D-xylulose 5 phosphate reductoisomerase.clm.	6
	L2	1-deoxy-D-xylulose 5 phosphate reductoisomerase	19
	DB=US	SPT; PLUR=YES; OP=ADJ	
	L1	6465225	1

END OF SEARCH HISTORY

h

=> file medline caplus biosis embase biotechds scisearch COST IN U.S. DOLLARS SINCE FILE

ENTRY 0.42 TOTAL SESSION 0.42

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 11:33:02 ON 13 JAN 2004

FILE 'CAPLUS' ENTERED AT 11:33:02 ON 13 JAN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS) Searches Updated 1/13/04

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FILE 'SCISEARCH' ENTERED AT 11:33:02 ON 13 JAN 2004 COPYRIGHT 2004 THOMSON ISI

=> 1-deoxy-D-xylulose 5 phosphate reductoisomerase 1-DEOXY-D-XYLULOSE IS NOT A RECOGNIZED COMMAND The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter "HELP COMMANDS" at an arrow prompt (=>).

=> s l1 and isoprenoid L2 162 L1 AND ISOPRENOID

=> s 12 and (dna or nucleic acid or rna)
2 FILES SEARCHED...

L3 26 L2 AND (DNA OR NUCLEIC ACID OR RNA)

=> dup rem 14
PROCESSING COMPLETED FOR L4
L5 10 DUP REM L4 (2 DUPLICATES REMOVED)

=> 15 and 1990-1998/py
L5 IS NOT A RECOGNIZED COMMAND
The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
"HELP COMMANDS" at an arrow prompt (=>).

=> s 15 and 1990-1998/py 4 FILES SEARCHED... L6 1 L5 AND 1990-1998/PY

=> d l6 ibib ab

L6 ANSWER 1 OF 1 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED. on STN

ACCESSION NUMBER:

1998305569 EMBASE A 1-deoxy-D-xylulose

5-phosphate reductoisomerase catalyzing the formation of 2-C-

methyl-D-erythritol 4

-phosphate in an alternative nonmevalonate

pathway for terpenoid biosynthesis.

AUTHOR:

Takahashi S.; Kuzuyama T.; Watanabe H.; Seto H.

CORPORATE SOURCE:

H. Seto, Inst. of Molec./Cellular Biosciences, University

of Tokyo, Bunkyo-ku, Tokyo 113-0032, Japan.

c00402@simail.nc.jp

SOURCE:

Proceedings of the National Academy of Sciences of the United States of America, (18 Aug 1998) 95/17 (9879-9884).

Refs: 38

ISSN: 0027-8424 CODEN: PNASA6

COUNTRY: DOCUMENT TYPE: United States Journal; Article Microbiology 004

FILE SEGMENT:

Clinical Biochemistry 029

LANGUAGE:

English

English SUMMARY LANGUAGE:

Several eubacteria including Esherichia coli use an alternative nonmevalonate pathway for the biosynthesis of isopentenyl diphosphate instead of the ubiquitous mevalonate pathway. In the alternative pathway, 2-C- methyl-D-erythritol or its 4-phosphate, which is proposed to be formed from 1-deoxy-D-xylulose 5-phosphate via intramolecular rearrangement followed by reduction process, is one of the biosynthetic precursors of isopentenyl diphosphate. To clone the gene(s) responsible for synthesis of 2-C-methyl-D-

erythritol 4-phosphate, we prepared and

selected E. coli mutants with an obligatory requirement for 2-C-methylerythritol for growth and survival. All the DNA fragments that complemented the defect in synthesizing 2-

C-methyl-D- erythritol 4-

phosphate of these mutants contained the yaeM gene, which is located at 4.2 min on the chromosomal map of E. coli. The gene product showed significant homologies to hypothetical proteins with unknown functions present in Haemophilus influenzae, Synechocystis sp. PCC6803, Mycobacterium tuberculosis, Helicobacter pyroli, and Bacillus subtilis. The purified recombinant yaeM gene product was overexpressed in E. coli and found to catalyze the formation of 2-C-

methyl-D-erythritol 4-

phosphate from 1-deoxy- D-xylulose 5-phosphate in the presence of NADPH. Replacement of NADPH with NADH decreased the reaction rate to about 1% of the original rate. The enzyme required Mn2+, Co2+, or Mg2+ as well. These data clearly show that the yaeM gene encodes an enzyme, designated 1-deoxy-D-xylulose 5-

phosphate reductoisomerase, that synthesizes 2

-C-methyl-D-erythritol 4

-phosphate from 1-deoxy-D-xylulose 5-phosphate, in a single step by intramolecular rearrangement and reduction and that this gene is responsible for terpenoid biosynthesis in E. coli.

=> d his

(FILE 'HOME' ENTERED AT 11:32:03 ON 13 JAN 2004)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, BIOTECHDS, SCISEARCH' ENTERED AT 11:33:02 ON 13 JAN 2004

297 S 1-DEOXY-D-XYLULOSE 5 PHOSPHATE REDUCTOISOMERASE T.1

162 S L1 AND ISOPRENOID L2

26 S L2 AND (DNA OR NUCLEIC ACID OR RNA) L3

12 S L3 AND 2-C METHYL-D-ERYTHRITOL 4-PHOSPHATE T<sub>1</sub>4

10 DUP REM L4 (2 DUPLICATES REMOVED)  $L_5$ 

1 S L5 AND 1990-1998/PY L6

ANSWER 1 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2002:123201 CAPLUS

DOCUMENT NUMBER:

136:162385

TITLE:

Methyl-D-erythritol phosphate pathway gene gdpE from

Arabidopsis thaliana and other plants

INVENTOR(S):

Boronat, Albert; Campos, Narciso; Rodriguez-

Concepcion, Manuel; Rohmer, Michel; Seeman, Myriam;

Valentin, Henry E.; Venkatesh, Tyamagondlu V.;

Venkatramesh, Mylavarapu

PATENT ASSIGNEE(S):

Monsanto Technology, LLC, USA

SOURCE:

PCT Int. Appl., 155 pp.

DOCUMENT TYPE:

Patent

CODEN: PIXXD2

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE	DATE										
WO 2002012478 A2 20020214 WO 2001-US24335 20010806	20010806										
WO 2002012478 C1 20020704											
WO 2002012478 A3 20030703											
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, Cl	H, CN,										
CO. CR. CU. CZ. DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, G	E, GH,										
GM. HR. HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, L	K, LR,										
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, P.	L, PT,										
RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, U	rG, UZ,										
VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM											
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, C	H, CY,										
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, T	R, BF,										
BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, T	'G										
AU 2001090522 A5 20020218 AU 2001-90522 20010806											
US 2002069426 A1 20020606 US 2001-921992 20010806											
EP 1356033 A2 20031029 EP 2001-970529 20010806											
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, M	IC, PT,										
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR											
PRIORITY APPLN. INFO.: US 2000-223483P P 20000807											
WO 2001-US24335 W 20010806											

The present invention provides and includes nucleic acids, proteins and AB antibodies assocd. with novel genes in the methyl-D-erythritol phosphate (MEP) biosynthesis pathway. Specifically, a homolog of the Escherichia coli gcpE gene is found in Arabidopsis thaliana which catalyzes the conversion of 2-C-methyl-D-erythritol 2,4-cyclodiphophate to (E)-1-(4-hydroxy-3-methylbut-2-enyl) diphosphate. Partial gene sequences are also provided from soybean, tomato, Mesembryanthemum crystallinum, rice, maize, loblolly pine, soybean, Brassica, and Physcomitrella patens. The invention further encompasses methods utilizing such mols., for example in gene isolation, gene anal. and the prodn. of transgenic plants. The present invention also includes transgenic plants modified to express proteins assocd. with the MEP pathway. Modulation of isoprenoid , tocopherol, monoterpene, and carotenoid levels can be achieved in transgenic plants.

ANSWER 2 OF 10

MEDLINE on STN

ACCESSION NUMBER:

2001489332 MEDLINE

DOCUMENT NUMBER:

21425086 PubMed ID: 11532167

TITLE:

1-Deoxy-D-xylulose

5-phosphate reductoisomerase

and plastid isoprenoid biosynthesis during tomato

fruit ripening.

AUTHOR:

Rodriguez-Concepcion M; Ahumada I; Diez-Juez E;

Sauret-Gueto S; Lois L M; Gallego F; Carretero-Paulet L;

Campos N; Boronat A

CORPORATE SOURCE:

Departament de Bioquimica i Biologia Molecular, Facultat de Quimica, Universitat de Barcelona, Marti i Franques 1-7,

08028 Barcelona, Spain.. mrodrigu@sun.bq.ub.es

PLANT JOURNAL, (2001 Aug) 27 (3) 213-22. SOURCE:

Journal code: 9207397. ISSN: 0960-7412.

PUB. COUNTRY:

England: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT: OTHER SOURCE: Priority Journals GENBANK-AF331705

ENTRY MONTH:

200112

ENTRY DATE:

Entered STN: 20010905

Last Updated on STN: 20020122 Entered Medline: 20011204

The recently discovered 2-C-methyl-D AΒ

-erythritol 4-phosphate (MEP) pathway for

the biosynthesis of plastid isoprenoids (including carotenoids) is not fully elucidated yet despite its central importance for plant life. It is known, however, that the first reaction completely specific to the pathway is the conversion of 1-deoxy-D-xylulose 5-phosphate (DXP) into MEP by the enzyme DXP reductoisomerase (DXR). We have identified a tomato cDNA encoding a protein with homology to DXR and in vivo activity, and show that the levels of the corresponding DXR mRNA and encoded protein in fruit tissues are similar before and during the massive accumulation of carotenoids characteristic of fruit ripening. The results are consistent with a non-limiting role of DXR, and support previous work proposing DXP synthase (DXS) as the first regulatory enzyme for plastid isoprenoid biosynthesis in tomato fruit. Inhibition of DXR activity by fosmidomycin showed that plastid isoprenoid biosynthesis is required for tomato fruit carotenogenesis but not for. other ripening processes. In addition, dormancy was reduced in seeds from fosmidomycin-treated fruit but not in seeds from the tomato yellow ripe mutant (defective in phytoene synthase-1, PSY1), suggesting that the isoform PSY2 might channel the production of carotenoids for abscisic acid biosynthesis. Furthermore, the complete arrest of tomato seedling development using fosmidomycin confirms a key role of the MEP pathway in plant development.

ANSWER 3 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2001:832956 CAPLUS

DOCUMENT NUMBER:

137:16268

TITLE:

First-time isolation of an isoprene synthase gene and heterologous expression of a gene originating from poplar and characterization of entry genes of a

mevalonate-independent isoprenoid

biosynthesis path from the cyanobacterium Synechoccus

leopoliensis Miller, Barbara

CORPORATE SOURCE:

Germany

SOURCE:

AUTHOR(S):

Schriftenreihe des Fraunhofer-Instituts

Atmosphaerische Umweltforschung (2001), 68,

a,b,c,d,e,f,i-v,1-145

CODEN: SFAUFS; ISSN: 1436-1094

Shaker Verlag

PUBLISHER: DOCUMENT TYPE:

Journal German

LANGUAGE:

The occurrence of 2-C-methyl-D-AB

erythritol-4-phosphate (MEP) metab. was

evidenced in cyanobacteria. A cosmid gene bank of Synechococcus leopoliensis contg. 1384 clones was provided. The genes encoding deoxyxylulosephosphate (DXP) synthase (DXS) and DXP reductoisomerase (DXR) were localized. Open reading frames of the dxs and dxr operons were identified, cloned, and expressed in Escherichia coli. The overexpression resulted in an 8-fold increase of the content of dimethylallyl diphosphate. The functionality of DXR was evidenced by photometric detn. of the NADPH oxidn. dependent on DXP. An isoprene synthase gene was isolated from a .lambda.-gene bank of the plant hybrid Populus alba x P.

tremula. High homologies to monoterpene synthases were found by sequencing. Overexpression of the isoprene synthase gene contg. an N-terminal signal peptide caused a 100-fold increase in isoprene formation. Simultaneous overexpression of the dxs gene of S. leopoliensis addnl. increased isoprene formation fourfold. Recombinant isoprene synthase increased the isoprene formation 200-fold in comparison with the limonene formation dependent on geranylphosphate.

REFERENCE COUNT:

THERE ARE 152 CITED REFERENCES AVAILABLE FOR 152 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 4 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN 1.5

ACCESSION NUMBER:

2001:158099 CAPLUS

DOCUMENT NUMBER:

134:291006

TITLE:

Investigation of a novel biosynthetic pathway to

isopentenyl diphosphate in Escherichia coli and Zymomonas mobilis: Identification and characterization

of involved genes

AUTHOR(S):

Grolle, Sigrid

CORPORATE SOURCE:

Germany

SOURCE:

Berichte des Forschungszentrums Juelich (2000),

Juel-3799, i-ix, 1-111

CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE:

Report

LANGUAGE:

German In the last few years, evidence has emerged that in bacteria a novel biosynthetic pathway to isopentenyl diphosphate exists. In this pathway isopentenyl diphosphate is formed from pyruvate and glyceraldehyde 3-phosphate. In the 1st reaction step these C3 compds. are combined to 1-deoxyxylulose 5-phosphate whereby pyruvate is decarboxylated. The gene encoding the 1-deoxyxylulose 5-phosphate synthase was identified in E. coli by searching for transketolase-homologous genes. The corresponding gene product was purified and was shown by NMR anal. to catalyze in a thiamin diphosphate (TPP) and Mg2+ dependent reaction the synthesis of 1-deoxyxylulose 5-phosphate. Phylogenetic investigation revealed that the 1-deoxyxylulose 5-phosphate synthase belongs to a new family of TPP dependent enzymes. The dxs gene is located at 9 min on the E. coli chromosome and is organized in 1 operon with a putative aldoketo-reductase gene. The disruption of this E. coli gene yielded no phenotype which indicates that the gene is not involved in the novel pathway. The gene encoding the second enzyme of the pathway, the 1-deoxyxylulose 5-phosphate reductoisomerase, was isolated from Z. mobilis. The 1-deoxyxylulose 5-phosphate reductoisomerase catalyzes the NADPH and Mn2+ dependent rearrangement and subsequent redn. of 1-deoxyxylulose 5-phosphate to 2C-methylerythritol 4-phosphate. The enzyme activity is competitively inhibited by the antibiotic fosmidomycin with a Ki of 0,6 .mu.M. By a E. coli strain engineered to produce the isoprenoid zeaxanthin the gene encoding IPP-isomerase, but no further genes of the novel pathway

REFERENCE COUNT:

could be isolated from an E. coli expression gene library. THERE ARE 236 CITED REFERENCES AVAILABLE FOR 236 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 5 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN

2000:34841 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

132:89232

TITLE:

Vitamin production by fermentative biosynthesis of

intermediates using genetically engineered

microorganisms followed by chemical synthesis

Millis, James R.; Saucy, Gabriel G.; Maurina-Brunker, INVENTOR(S): Julie; McMullin, Thomas W.

DCV, Inc., USA PATENT ASSIGNEE(S):

SOURCE:

PCT Int. Appl., 239 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

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APPLICATION NO. DATE
    PATENT NO.
                   KIND DATE
                                            _____
     ______
    WO 2000001650 A1
                            20000113 WO 1999-US15264 19990706
        W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
             DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
            JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                            AU 1999-48630
                       A1
                            20000124
     AU 9948630
                                           EP 1999-932295
                                                              19990706
                       Α1
                             20010502
    EP 1095002
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                            US 1999-348097
                                                              19990706
     US 6410755
                       B1
                            20020625
                                            JP 2000-558056 19990706
                       T2
                             20020702
     JP 2002519049
                                            US 1999-350275
                                                              19990706
                       B1
                             20030311
     US 6531303
                                            US 2001-902187
                                                              20010709
     US 2003125573
                       A1
                             20030703
                    A1
                                           US 2001-909558
                                                              20010720
     US 2003092144
                             20030515
                                         US 1998-91951P P 19980706
PRIORITY APPLN. INFO.:
                                         US 1998-91964P P 19980706
                                         US 1998-91983P P 19980706
                                         US 1998-91868P P 19980706
                                         US 1999-348097 A1 19990706
                                         US 1999-350275 A1 19990706
                                         WO 1999-US15264 W 19990706
```

OTHER SOURCE(S): MARPAT 132:89232

The invention provides methods of producing vitamin E (.alpha.-tocopherol and .alpha.-tocopheryl esters), vitamin A (retinol), or .beta.-carotene. The methods comprise using a biol. system to produce farnesol or geranylgeraniol. Biosynthesis of the farnesol or geranylgeraniol intermediates is enhanced by shifting microbial metab. away from sterol biosynthesis via genetic inactivation of the squalene synthase ERG9 gene or by inactivation of squalene synthase by zaragozic acid in a strain with a functional ERG9 gene. Geranylgeraniol biosynthesis is further enhanced in strains over-expressing any of 4 different cloned geranylgeranyl pyrophosphate synthase genes: (1) BTS1 gene from Saccharomyces cerevisiae; (2) crtE gene from Erwinia uredovora; (3) a1-3 gene from Neurospora crassa; or (4) ggs gene from Gibberella fujikuroi. Overexpressing of hydroxymethyl-CoA reductase and/or the ERG20 gene which encodes farnesyl pyrophosphate synthase in Saccharomyces cerevisiae also enhances biosynthesis of fermentative intermediates. Finally, over-expression of multiple isoprenoid pathway genes or alternative pathway (Rohmer pathway) was further investigated in strains that have an erg9 mutation and elevated levels of hydroxymethylglutaryl-CoA reductase. The farnesol or geranylgeraniol fermn. products are then chem. converted into .alpha.-tocopherol, an .alpha.-tocopheryl ester, vitamin A, or .beta. carotene.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 10 SCISEARCH COPYRIGHT 2004 THOMSON ISI ON STN ACCESSION NUMBER: 2000:516559 SCISEARCH
THE GENUINE ARTICLE: 330AU

3

TITLE:

Characterization of 1-Deoxy-D

-xylulose 5-phosphate

reductoisomerase, an enzyme involved in isopentenyl diphosphate biosynthesis, and identification of its catalytic amino acid residues

AUTHOR:

Kuzuyama T; Takahashi S; Takagi M; Seto H (Reprint)

CORPORATE SOURCE:

UNIV TOKYO, INST MOL & CELLULAR BIOSCI, BUNKYO KU, TOKYO 1130032, JAPAN (Reprint); UNIV TOKYO, INST MOL & CELLULAR

BIOSCI, BUNKYO KU, TOKYO 1130032, JAPAN

COUNTRY OF AUTHOR:

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (30 JUN 2000) Vol. 275,

No. 26, pp. 19928-19932.

Publisher: AMER SOC BIOCHEMISTRY MOLECULAR BIOLOGY INC,

9650 ROCKVILLE PIKE, BETHESDA, MD 20814.

ISSN: 0021-9258.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT: LANGUAGE:

LIFE English

REFERENCE COUNT:

26

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

1-Deoxy-D-xylulose 5-phosphate (DXP) reductoisomerase, which AB simultaneously catalyzes the intramolecular rearrangement and reduction of

DXP to form 2-C-methyl-Derythritol 4-phosphate, constitutes a key

enzyme of an alternative mevalonate-independent pathway for isopentenyl diphosphate biosynthesis. The dxr gene encoding this enzyme from Escherichia coli was overexpressed as a histidine-tagged protein and characterized in detail. DNA sequencing analysis of the dxr genes from 10 E. coli dxr-deficient mutants revealed base substitution mutations at four points: two nonsense mutations and two amino acid substitutions (Gly(14) to Asp(14) and Glu(231) to Lys(231)), Diethyl pyrocarbonate treatment inactivated DXP reductoisomerase, and subsequent hydroxylamine treatment restored the activity of the diethyl pyrocarbonate-treated enzyme. To characterize these defects, we overexpressed the mutant enzymes G14D, E231K, H153Q, H209Q, and H257Q. All of these mutant enzymes except for G14D were obtained as soluble proteins. Although the purified enzyme E231K had wildtype K-m values for DXP and NADPH, the mutant enzyme had less than a 0.24% wild-type k(cat) value. K-m values of H153Q, H209Q, and H257Q for DXP increased to 3.5-, 7.6-, and 19-fold the wild-type value, respectively. These results indicate that Glu(231) of E. coli DXP reductoisomerase plays an important role(s) in the conversion of DXP to 2-C-methyl-D-

erythritol 4-phosphate, and that His(153), His(209), and His(257), in part, associate with DXP binding in the enzyme molecule.

ANSWER 7 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN L5

2000:140254 CAPLUS ACCESSION NUMBER:

132:275501

DOCUMENT NUMBER: TITLE:

SOURCE:

Overlooked nonmevalonate pathway for isopentenyl

diphosphate biosynthesis and specific inhibitors

AUTHOR(S):

Kuzuyama, Tomohisa; Seto, Haruo

CORPORATE SOURCE:

Institute of Molecular and Cellular Biosciences,

University of Tokyo, Tokyo, 113-0032, Japan Nihon Yukagakkaishi (2000), 49(2), 119-125

CODEN: NIYUFC; ISSN: 1341-8327

PUBLISHER: DOCUMENT TYPE:

Nihon Yukagaku Gakkai

Journal; General Review

LANGUAGE:

Japanese

A review with 19 refs. Several eubacteria including Escherichia coli utilize a mevalonate-independent pathway (nonmevalonate pathway) for the biosynthesis of isopentenyl diphosphate. In the nonmevalonate pathway, 2-C-methyl-D-erythritol or its 4-phosphate, possibly formed from 1-deoxy-D-xylulose 5-phosphate (DXP) via intramol. rearrangement followed by redn., is a biosynthetic precursors of isopentenyl diphosphate. To clone the gene(s) responsible for the synthesis of 2-C -methyl-D-erythritol 4-

phosphate (MEP), E. coli mutants with obligatory requirement for 2-C-methylerythritol for growth and survival were prepd. DNA fragments complementing the defect in synthesizing MEP of these mutants contained the yaeM gene located at 4.2 min on the chromosomal map of E. coli. The gene product showed significant homol. to hypothetical proteins with unknown functions in many eubacteria. The yaeM gene product overexpressed in E. coli was found to catalyze the formation of MEP from DXP in the presence of NADPH. The yaeM gene is thus clearly shown to encode a novel enzyme, DXP reductoisomerase, which synthesizes MEP from DXP in a single step by intramol. rearrangement and redn. Fosmidomycin was noted to be a specific inhibitor of DXP reductoisomerase.

L5 ANSWER 8 OF 10 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

DUPLICATE 1

ACCESSION NUMBER: 1999:247659 BIOSIS DOCUMENT NUMBER: PREV199900247659

TITLE:

Isoprenoid biosynthesis via a

mevalonate-independent pathway in plants: Cloning and

heterologous expression of 1-deoxy-

D-xylulose-5-phosphate

reductoisomerase from peppermint.

AUTHOR(S):
CORPORATE SOURCE:

Lange, B. Markus; Croteau, Rodney [Reprint author]
Department of Biochemistry and Biophysics, Institute of
Biological Chemistry, Washington State University, Pullman,

WA, 99164-6340, USA

SOURCE:

Archives of Biochemistry and Biophysics, (May 1, 1999) Vol.

365, No. 1, pp. 170-174. print. CODEN: ABBIA4. ISSN: 0003-9861.

DOCUMENT TYPE:

Article English

LANGUAGE: ENTRY DATE:

Entered STN: 2 Jul 1999

Last Updated on STN: 2 Jul 1999

Two distinct pathways are utilized by plants for the biosynthesis of isopentenyl diphosphate, the universal precursor of isoprenoids. The classical acetate/mevalonate pathway operates in the cytosol, whereas plastidial isoprenoids originate via a novel mevalonate-independent route that involves a transketolase-catalyzed condensation of pyruvate and D-glyceraldehyde-3-phosphate to yield 1-deoxy-D-xylulose-5-phosphate as the first intermediate. Based on in vivo feeding experiments, rearrangement and reduction of deoxyxylulose phosphate have been proposed to give rise to 2-C-methyl-D-

erythritol-4-phosphate as the second

intermediate of this pyruvate/glyceraldehyde-3-phosphate pathway (1-3). The cloning of an Escherichia coli gene encoding an enzyme capable of converting 1-deoxy-D-xylulose-5-phosphate to 2-C-erythritol-4-phosphate was recently reported (4). A cloning strategy was developed for isolating the gene encoding a plant homolog of this enzyme from peppermint (Mentha X piperita), and the identity of the resulting cDNA was confirmed by heterologous expression in E. coli. Unlike the microbial reductoisomerase, the plant ortholog encodes a preprotein bearing an N-terminal plastidial transit peptide that directs the enzyme to plastids where the mevalonate-independent pathway operates in plants. The peppermint gene comprises an open reading frame of 1425 nucleotides which, when the plastidial targeting sequence is excluded, encodes a deduced enzyme of approximately 400 amino acid residues with a mature size of about 43.5 kDa.

L5 ANSWER 9 OF 10 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER: 1999241876 EMBASE

TITLE:

Cloning and heterologous expression of a cDNA encoding

1-deoxy-D-xylulose-

5-phosphate reductoisomerase of

Arabidopsis thaliana.

AUTHOR:

CORPORATE SOURCE:

Schwender J.; Muller C.; Zeidler J.; Lichtenthaler H.K. H.K. Lichtenthaler, Botanisches Institut, Universitat

Karlsruhe, D-76128 Karlsruhe, Germany.

hartmut.lichtenthaler@bio-geo.uni-karlsruhe.de

SOURCE: FEBS Letters, (1999) 455/1-2 (140-144).

Refs: 22

ISSN: 0014-5793 CODEN: FEBLAL

PUBLISHER IDENT.: S 0014-5793 (99) 00849-2

COUNTRY: Netherlands
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 029 Clinical Biochemistry

LANGUAGE: English SUMMARY LANGUAGE: English

AB Various plant isoprenoids are synthesized via the non-mevalonate pathway of isopentenyl diphosphate formation. In this pathway, 1-deoxy-D-xylulose 5-phosphate (DOXP), the first intermediate, is transformed to 2-

C-methyl-D-erythritol 4-

phosphate (MEP) by an enzyme which was recently cloned from
Escherichia coli. In order to find a plant homologue of this 1-

deoxy-D-xylulose 5-phosphate

reductoisomerase (DXR) we cloned a cDNA fragment from Arabidopsis thaliana which has high homology to the E. coli DXR. By expression of this fragment in E. coli we could demonstrate that it encodes a protein which transforms DOXP to MEP. The antibiotic fosmidomycin specifically inhibits this DXR enzyme activity. Copyright (C) 1999 Federation of European Biochemical Societies.

L5 ANSWER 10 OF 10 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN DUPLICATE 2

ACCESSION NUMBER: 1998305569 EMBASE
TITLE: A 1-deoxy-D-xylulose

5-phosphate reductoisomerase catalyzing the formation of 2-C-

methyl-D-erythritol 4

-phosphate in an alternative nonmevalonate

pathway for terpenoid biosynthesis.

AUTHOR: Takahashi S.; Kuzuyama T.; Watanabe H.; Seto H.

CORPORATE SOURCE: H. Seto, Inst. of Molec./Cellular Biosciences, University

of Tokyo, Bunkyo-ku, Tokyo 113-0032, Japan.

c00402@simail.nc.jp

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America, (18 Aug 1998) 95/17 (9879-9884).

Refs: 38

ISSN: 0027-8424 CODEN: PNASA6

COUNTRY: United States
DOCUMENT TYPE: Journal; Article
FILE SEGMENT: 004 Microbiology

029 Clinical Biochemistry

LANGUAGE: English SUMMARY LANGUAGE: English

AB Several eubacteria including Esherichia coli use an alternative nonmevalonate pathway for the biosynthesis of isopentenyl diphosphate instead of the ubiquitous mevalonate pathway. In the alternative pathway, 2-C- methyl-D-erythritol or its 4-phosphate, which is proposed to be formed from 1-deoxy-D-xylulose 5-phosphate via intramolecular rearrangement followed by reduction process, is one of the biosynthetic precursors of isopentenyl diphosphate. To clone the gene(s) responsible for synthesis of 2-C-methyl-D-

erythritol 4-phosphate, we prepared and

selected E. coli mutants with an obligatory requirement for 2-C-methylerythritol for growth and survival. All the DNA fragments that complemented the defect in synthesizing 2-

C-methyl-D- erythritol 4-

phosphate of these mutants contained the yaeM gene, which is located at 4.2 min on the chromosomal map of E. coli. The gene product showed significant homologies to hypothetical proteins with unknown functions present in Haemophilus influenzae, Synechocystis sp. PCC6803, Mycobacterium tuberculosis, Helicobacter pyroli, and Bacillus subtilis. The purified recombinant yaeM gene product was overexpressed in E. coli

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and found to catalyze the formation of 2-C-
    methyl-D-erythritol 4-
    phosphate from 1-deoxy- D-xylulose 5-phosphate in the presence of
    NADPH. Replacement of NADPH with NADH decreased the reaction rate to about
     1% of the original rate. The enzyme required Mn2+, Co2+, or Mg2+ as well.
    These data clearly show that the yaeM gene encodes an enzyme, designated
     1-deoxy-D-xylulose 5-
    phosphate reductoisomerase, that synthesizes 2
     -C-methyl-D-erythritol 4
     -phosphate from 1-deoxy-D-xylulose 5-phosphate, in a single step
     by intramolecular rearrangement and reduction and that this gene is
     responsible for terpenoid biosynthesis in E. coli.
=> d his
     (FILE 'HOME' ENTERED AT 11:32:03 ON 13 JAN 2004)
     FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, BIOTECHDS, SCISEARCH' ENTERED AT
     11:33:02 ON 13 JAN 2004
            297 S 1-DEOXY-D-XYLULOSE 5 PHOSPHATE REDUCTOISOMERASE
            162 S L1 AND ISOPRENOID
             26 S L2 AND (DNA OR NUCLEIC ACID OR RNA)
             12 S L3 AND 2-C METHYL-D-ERYTHRITOL 4-PHOSPHATE
             10 DUP REM L4 (2 DUPLICATES REMOVED)
              1 S L5 AND 1990-1998/PY
=> dup rem 13
PROCESSING COMPLETED FOR L3
             20 DUP REM L3 (6 DUPLICATES REMOVED)
=> s 17 and 1993-1998/py
             1 L7 AND 1993-1998/PY
=> d 18
     ANSWER 1 OF 1 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
     1998305569 EMBASE
     A 1-deoxy-D-xylulose 5-
     phosphate reductoisomerase catalyzing the formation of
     2-C-methyl-D-erythritol 4-phosphate in an alternative nonmevalonate
     pathway for terpenoid biosynthesis.
     Takahashi S.; Kuzuyama T.; Watanabe H.; Seto H.
     H. Seto, Inst. of Molec./Cellular Biosciences, University of Tokyo,
     Bunkyo-ku, Tokyo 113-0032, Japan. c00402@simail.nc.jp
     Proceedings of the National Academy of Sciences of the United States of
     America, (18 Aug 1998) 95/17 (9879-9884).
     Refs: 38
     ISSN: 0027-8424 CODEN: PNASA6
     United States
     Journal; Article
     004
             Microbiology
             Clinical Biochemistry
     029
     English
     English
=> s 1-deoxy-D-xylulose 5 phosphate reductoisomerase and e. coli
            69 1-DEOXY-D-XYLULOSE 5 PHOSPHATE REDUCTOISOMERASE AND E. COLI
=> dup rem 19
PROCESSING COMPLETED FOR L9
             38 DUP REM L9 (31 DUPLICATES REMOVED)
L10
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L1

L2

L3

L4L5

1.6

 $L_8$ 

1.8

ΑN

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CS

SO

CY

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LA

SL

L9

=> focus 110

PROCESSING COMPLETED FOR L10 38 FOCUS L10 1-

=> d l11 1-5 ibib ab

L11 ANSWER 1 OF 38 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2001:780314 CAPLUS

DOCUMENT NUMBER:

135:340826

TITLE:

Method for the determination of 1-

deoxy-D-xylulose 5

-phosphate reductoisomerase in microorganisms and cell cultures

INVENTOR(S):

Bacher, Adelbert; Eisenreich, Wolfgang; Fellermeier, Monika; Hecht, Stefan; Herz, Stefan; Rohdich, Felix;

Wungsintaweekul, Juraithip; Zenk, Meinhart H.

PATENT ASSIGNEE(S):

Germany

SOURCE:

Ger. Offen., 8 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE

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APPLICATION NO.

DE 10018368 A1

20011025

\_\_\_\_\_

DE 2000-10018368 20000413

\_\_\_\_\_ PRIORITY APPLN. INFO.:

DE 2000-10018368

\_\_\_\_\_\_

The invention concerns an assay for the detn. of 1-deoxy AB

-D-xylulose 5-phosphate

reductoisomerase in microorganisms and plant cell cultures by using 1-deoxy-D-xylulose as substrate and a phosphorylation agent in the presence of a magnesium salt, sodium fluoride and glutathione. 1

-Deoxy-D-xylulose 5-

phosphate reductoisomerase is detd. in genetically engineered E.coli; radiolabeled substrate can be used. Thus a reagent contained 50 mM Tris-HCl pH 7.4, 40 mM MgCl2, 40 mM ATP, 20 mM glutathione, 20 mM NaF, and 3.5 .mu.M [1,2-14C]1-deoxy-D-xylulose (24000 dpm) in 50 .mu.L including the sample. After incubation at 37.degree.C for 1 h paper chromatog. was performed; Rf values were detd. with a radioactivity reader.

L11 ANSWER 2 OF 38 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2000:772741 CAPLUS

DOCUMENT NUMBER:

133:330553

TITLE:

Arabidopsis 1-deoxy-Dxylulose-5-phosphate

reductoisomerase cDNA and transgenic plants

with enhanced tocopherol content

INVENTOR(S):

Lichtenthaler, Hartmut; Schwender, Jorg; Reindl,

Andreas; Herbers, Karin

PATENT ASSIGNEE(S): SOURCE:

BASF Aktiengesellschaft, Germany

PCT Int. Appl., 41 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2000065036

A2 20001102 WO 2000-EP3465 20000417

WO 2000065036

A3 20010419

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,

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ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                             DE 1999-19918949 19990427
                             20001123
     DE 19918949
                        A1
     EP 1180149
                             20020220
                                             EP 2000-922642
                                                                20000417
                        A2
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                          DE 1999-19918949 A
                                                                19990427
PRIORITY APPLN. INFO .:
                                          WO 2000-EP3465
                                                            W
                                                                20000417
     The invention relates to a method for producing plants contg. increased
     quantities of tocopherols, vitamin K, carotenoids, chlorophylls and
     polyterpenes by overexpression of a 1-deoxy-D
     -xylulose-5-phosphate
     reductoisomerase (DXPRI) gene. Thus, the DXPRI cDNA of
     Arabidopsis thaliana was cloned, sequenced, and expressed in E.
     coli, tobacco, and Brassica napus. The .alpha.-tocopherol levels
     of the transgenic plants were increased.
L11 ANSWER 3 OF 38 CAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:
                          1998:358885 CAPLUS
DOCUMENT NUMBER:
                          129:149156
                          Direct formation of 2-C-methyl-D-erythritol
TITLE:
                          4-phosphate from 1-deoxy-D-xylulose 5-phosphate by
                           1-deoxy-D-xylulose
                           5-phosphate reductoisomerase
                             a new enzyme in the non-mevalonate pathway to
                           isopentenyl diphosphate
                           Kuzuyama, Tomohisa; Takahashi, Shunji; Watanabe,
AUTHOR (S):
                          Hiroyuki; Seto, Haruo
                           Institute of Molecular and Cellular Biosciences,
CORPORATE SOURCE:
                          University of Tokyo, Tokyo, 113-0032, Japan
                          Tetrahedron Letters (1998), 39(25), 4509-4512
SOURCE:
                          CODEN: TELEAY; ISSN: 0040-4039
                          Elsevier Science Ltd.
PUBLISHER:
DOCUMENT TYPE:
                          Journal
                           English
LANGUAGE:
     1-Deoxy-D-xylulose 5-phosphate is biotransformed to 2-C-methyl-D-
     erythritol 4-phosphate in a single step in the presence of NADPH by a new
     recombinant enzyme named 1-deoxy-D-
     xylulose 5-phosphate reductoisomerase
     purified from Escherichia coli.
                                 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                           19
                                 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L11 ANSWER 4 OF 38 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN
                     2000:95738 BIOSIS
ACCESSION NUMBER:
                     PREV200000095738
DOCUMENT NUMBER:
                     Biosynthesis of terpenoids: 1-Deoxy-
TITLE:
                     D-xylulose-5-phosphate
                     reductoisomerase from Escherichia coli is a class B
                     dehydrogenase.
                     Radykewicz, Tanja; Rohdich, Felix; Wungsintaweekul,
AUTHOR (S):
                     Juraithip; Herz, Stefan; Kis, Klaus; Eisenreich, Wolfgang;
                     Bacher, Adelbert; Zenk, Meinhart H.; Arigoni, Duilio
                      [Reprint author]
                     Laboratorium fur Organische Chemie, ETH Zurich,
CORPORATE SOURCE:
                     Universitatsstr. 16, CH-8092, Zurich, Switzerland
                     FEBS Letters, (Jan. 14, 1999) Vol. 464, No. 2-3, pp.
SOURCE:
```

Article DOCUMENT TYPE:

157-160. print.

CODEN: FEBLAL. ISSN: 0014-5793.

AΒ

LANGUAGE:

English

ENTRY DATE:

Entered STN: 15 Mar 2000

Last Updated on STN: 3 Jan 2002

1-Deoxy-D-xylulose-5-phosphate is converted into 2-C-methyl-D-erythritol-4phosphate by the catalytic action of 1-deoxy-D

-xylulose-5-phosphate

reductoisomerase (Dxr protein) using NADPH as cofactor. The stereochemical features of this reaction were investigated in in vitro experiments with the recombinant Dxr protein of Escherichia coli using (4R) - or (4S) - (4-2H1) NADPH as coenzyme. The enzymatically formed 2-C-methyl-D-erythritol-4-phosphate was isolated and converted into 1,2:3,4-di-O-isopropylidene-2-C-methyl-D-erythritol; NMR spectroscopic investigation of this derivative indicated that only (4S)-(4-2H1)NADPH affords 2-C-methyl-D-erythritol-4-phosphate labelled exclusively in the HRe position of C-1. Stereospecific transfer of HSi from C-4 of the cofactor identifies the Dxr protein of E. coli as a class B dehydrogenase.

L11 ANSWER 5 OF 38 MEDLINE on STN ACCESSION NUMBER:

DOCUMENT NUMBER: 98374274

1998374274 MEDLINE

PubMed ID: 9707569

TITLE:

A 1-deoxy-D-xylulose

5-phosphate reductoisomerase

catalyzing the formation of 2-C-methyl-D-erythritol 4-phosphate in an alternative nonmevalonate pathway for

terpenoid biosynthesis.

AUTHOR:

Takahashi S; Kuzuyama T; Watanabe H; Seto H

CORPORATE SOURCE:

Institute of Molecular and Cellular Biosciences, University

of Tokyo, Bunkyo-ku, Tokyo 113-0032, Japan.

SOURCE:

PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (1998 Aug 18) 95 (17) 9879-84.

Journal code: 7505876. ISSN: 0027-8424.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English FILE SEGMENT:

OTHER SOURCE:

Priority Journals GENBANK-AB013300

ENTRY MONTH:

199809

ENTRY DATE:

Entered STN: 19980925

Last Updated on STN: 20030124 Entered Medline: 19980917

Several eubacteria including Esherichia coli use an alternative nonmevalonate pathway for the biosynthesis of isopentenyl diphosphate instead of the ubiquitous mevalonate pathway. In the alternative pathway, 2-C-methyl-D-erythritol or its 4-phosphate, which is proposed to be formed from 1-deoxy-D-xylulose 5-phosphate via intramolecular rearrangement followed by reduction process, is one of the biosynthetic precursors of isopentenyl diphosphate. To clone the gene(s) responsible for synthesis of 2-C-methyl-D-erythritol 4-phosphate, we prepared and selected E . coli mutants with an obligatory requirement for 2-C-methylerythritol for growth and survival. All the DNA fragments that complemented the defect in synthesizing 2-C-methyl-D-erythritol 4-phosphate of these mutants contained the yaeM gene, which is located at 4.2 min on the chromosomal map of E. coli. The gene product showed significant homologies to hypothetical proteins with unknown functions present in Haemophilus influenzae, Synechocystis sp. PCC6803, Mycobacterium tuberculosis, Helicobacter pyroli, and Bacillus subtilis. The purified recombinant yaeM gene product was overexpressed in E. coli and found to catalyze the formation of 2-C-methyl-D-erythritol 4-phosphate from 1-deoxy-D-xylulose 5-phosphate in the presence of NADPH. Replacement of NADPH with NADH decreased the reaction rate to about 1% of the original rate. The enzyme required Mn2+, Co2+, or Mg2+ as well. These data clearly show that the yaeM gene encodes

xylulose 5-phosphate reductoisomerase

an enzyme, designated 1-deoxy-D-

, that synthesizes 2-C-methyl-D-erythritol 4-phosphate from 1-deoxy-D-xylulose 5-phosphate, in a single step by intramolecular rearrangement and reduction and that this gene is responsible for terpenoid biosynthesis in **E. coli**.

#### => d l11 6-10 ibib ab

L11 ANSWER 6 OF 38 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1999:673053 CAPLUS

DOCUMENT NUMBER:

131:309853

TITLE:

Process for producing isoprenoid compounds by transgenic microorganisms and method for detecting compounds having antibacterial or herbicidal activity

INVENTOR(S):

Miyake, Koichiro; Hashimoto, Shinichi; Motoyama, Hiroaki; Ozaki, Akio; Seto, Haruo; Kuzuyama, Tomohisa;

Takahashi, Shunji

PATENT ASSIGNEE(S):

Kyowa Hakko Kogyo Co., Ltd., Japan

SOURCE:

PCT Int. Appl., 145 pp.

DOCUMENT TYPE:

CODEN: PIXXD2
Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.			KIND DATE				APPLICATION NO. DATE										
WO	9953 W:	AU,	BG,	BR,	CA,	CN,	CZ,	HU,	ID,	IL,	IN,	KR,	MX,	NO,	NZ,	PL,	
		ТJ,	$\mathbf{TM}$											KG,			
	RW:	AT, PT,		CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,
JP	2000300256			A2 20001031			JP 1999-104589						19990412				
JP	2000	2000300257			2	2000	1031		JP 1999-104590 19990412								
CA	2325	798		ΑZ	Ą	1999	1021		C	A 19	99-2	3257	98	1999	0414		
	9931				_	1999								1999	0414		
EP	1072													1999			
	R:	AT, IE,	BE, FI	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
PRIORIT	RIORITY APPLN. INFO			. :					JP 1	998-	1031	01	Α	1998	0414		
									JP 1	998-	2219	10	Α	1998	0805		
									JP 1	999-	3573	9	Α	1999	0215		
									WO 1	999-	JP19	87	W	1999	0414		

Disclosed is a method for the prodn. of isoprenoid compds. by cultivating transgenic prokaryotes that have been transformed with the gene for (1) 1-deoxy-D-xylulose 5-phosphate synthetase; (2) farnesyl pyrophosphate synthetase; (3) exodeoxyribo-nuclease; (4) a defined protein; and/or (5) 1-deoxy-D-xylulose 5-

phosphate reductoisomerase. The transgenic prokaryotes are selected from Escherichia, Rhodobacter, or Erwinia. The isoprenoid compds. are useful for (1) the treatment of heart diseases or osteoporosis, hemostasis, prevention of cancer, immunopotentiation, etc. and (2) the prepn. of health foods, antifouling coatings, etc. A method for screening compds. for their antibacterial or herbicidal activity or herbicidal activity by detecting their inhibitory activity against the enzymes assocd. with the non-mevalonate pathway is also claimed. Isolation of the genes assocd. with the biosynthesis of isoprenoid compds. from Escherichia coli strain XL1-Blue and use of the genes to improve the yield of CoQ8 by transgenic E. coli DH5.alpha. were shown. The Rhodobacter sphaeroides counterparts of gene DXS were also provided.

REFERENCE COUNT:

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 7 OF 38 MEDLINE on STN ACCESSION NUMBER: 2000123893 MEDLINE

DOCUMENT NUMBER: 20123893 PubMed ID: 10631325

TITLE: Biosynthesis of terpenoids: 1-deoxy-

D-xylulose-5-phosphate

reductoisomerase from Escherichia coli is a class B

dehydrogenase.

AUTHOR: Radykewicz T; Rohdich F; Wungsintaweekul J; Herz S; Kis K;

Eisenreich W; Bacher A; Zenk M H; Arigoni D

CORPORATE SOURCE: Lehrstuhl fur Organische Chemie und Biochemie, Technische

Universitat Munchen, Lichtenbergstr. 4, D-85747, Garching,

Germany.

SOURCE: FEBS LETTERS, (2000 Jan 14) 465 (2-3) 157-60.

Journal code: 0155157. ISSN: 0014-5793.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200002

ENTRY DATE: Entered STN: 20000309

Last Updated on STN: 20000309 Entered Medline: 20000218

AB 1-Deoxy-D-xylulose-5-phosphate is converted into 2-C-methyl-D-erythritol-4-

phosphate by the catalytic action of 1-deoxy-D

-xylulose-5-phosphate

reductoisomerase (Dxr protein) using NADPH as cofactor. The stereochemical features of this reaction were investigated in in vitro experiments with the recombinant Dxr protein of Escherichia coli using (4R) - or (4S) - [4 - (2) H(1)] NADPH as coenzyme. The enzymatically formed 2-C-methyl-D-erythritol-4-phosphate was isolated and converted into 1,2:3,4-di-O-isopropylidene-2-C-methyl-D-erythritol; NMR spectroscopic investigation of this derivative indicated that only (4S) - [4 - (2) H(1)] NADPH affords 2-C-methyl-D-erythritol-4-phosphate labelled exclusively in the H(Re) position of C-1. Stereospecific transfer of H(Si) from C-4 of the cofactor identifies the Dxr protein of E. coli as a class B dehydrogenase.

L11 ANSWER 8 OF 38 MEDLINE on STN

ACCESSION NUMBER: 1999355442 MEDLINE

DOCUMENT NUMBER: 99355442 PubMed ID: 10428488

TITLE: Cloning and heterologous expression of a cDNA encoding

1-deoxy-D-xylulose-

5-phosphate reductoisomerase of

Arabidopsis thaliana.

AUTHOR: Schwender J; Muller C; Zeidler J; Lichtenthaler H K

CORPORATE SOURCE: Botanisches Institut, Universitat Karlsruhe, Germany.

SOURCE: FEBS LETTERS, (1999 Jul 16) 455 (1-2) 140-4.
Journal code: 0155157. ISSN: 0014-5793.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals OTHER SOURCE: GENBANK-AJ242588

ENTRY MONTH: 199908

ENTRY DATE: Entered STN: 19990827

Last Updated on STN: 19990827 Entered Medline: 19990816

AB Various plant isoprenoids are synthesized via the non-mevalonate pathway of isopentenyl diphosphate formation. In this pathway, 1-deoxy-D-xylulose 5-phosphate (DOXP), the first intermediate, is transformed to 2-C-methyl-D-erythritol 4-phosphate (MEP) by an enzyme which was recently cloned from Escherichia coli. In order to find a plant homologue of this

1-deoxy-D-xylulose 5-phosphate reductoisomerase (DXR) we cloned a cDNA

fragment from Arabidopsis thaliana which has high homology to the

E. coli DXR. By expression of this fragment in

**E. coli** we could demonstrate that it encodes a protein which transforms DOXP to MEP. The antibiotic fosmidomycin specifically inhibits this DXR enzyme activity.

L11 ANSWER 9 OF 38 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2003:23430 CAPLUS

DOCUMENT NUMBER:

138:69471

TITLE:

Docking a ligand to a macromolecule by a combination of NMR measurements and computational modeling, and applications to protein-ligand interactions and

structure-based drug design

INVENTOR(S):

Sem, Daniel S.; Pellecchia, Maurizio

PATENT ASSIGNEE(S):

USA

SOURCE:

U.S. Pat. Appl. Publ., 32 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

US 2003008326 A1 20030109 US 2002-158770 20020530

US 2001-294675P P 20010530

US 2001-294675P P 20010530

PRIORITY APPLN. INFO : The present invention relates generally to interactions between macromols. and ligands and more specifically to NMR methods for detg. structure-related properties of a ligand when bound to a macromol. The invention provides a method for detg. a structure model for a test ligand bound to a macromol. binding site by a combination of NMR measurements and computational modeling. Structural constraints for the test ligand are derived from spectroscopic signals arising from interactions between the test ligand and macromol. The structure constraints are used as constraints in docking a structure model of the ligand to a structure model of the macromol., or as constraints in overlaying a structure model of the test ligand on the known structure for a ref. ligand that binds to the macromol. The invention further provides a method for detg. a structure model for a macromol. bound to a ligand. Structural constraints derived from spectroscopically obsd. interactions of the macromol. and a ref. ligand are used to guide mol. modeling or to evaluate the results of a mol. modeling simulation of the macromol. An advantage of the invention is that a structure model of a test ligand bound to the macromol. can be obtained at sufficient resoln. to assist in structure-based design of a biol. active agent or drug without the requirement for a complete detn. of the structure of the macromol.-test ligand complex. Examples include: docking of a furoic acid-based inhibitor into the NADH binding site of E. coli dihydrodipicolinate reductase (DHPR); overlay of a furoic acid-based inhibitor onto DHPR-bound NADH; validation of a binding site homol. model for 1-deoxy-Dxylulose-5-phosphate reductoisomerase

(DOXPR), and identifying a residue of DOXPR that is at an interface between ligand binding sites.

L11 ANSWER 10 OF 38 MEDLINE on STN

ACCESSION NUMBER: 2000387137

000387137 MEDLINE

DOCUMENT NUMBER:

20347905 PubMed ID: 10787409

TITLE:

Characterization of 1-deoxy-D

-xylulose 5-phosphate

reductoisomerase, an enzyme involved in isopentenyl diphosphate biosynthesis, and identification of its

catalytic amino acid residues.

AUTHOR:

Kuzuyama T; Takahashi S; Takagi M; Seto H

CORPORATE SOURCE:

Institute of Molecular and Cellular Biosciences, University

of Tokyo, Yayoi, Bunkyo-ku, Tokyo 113-0032, Japan.

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (2000 Jun 30) 275 (26)

19928-32.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200008

ENTRY DATE:

Entered STN: 20000818

Last Updated on STN: 20000818

Entered Medline: 20000810

AΒ 1-Deoxy-d-xylulose 5-phosphate (DXP) reductoisomerase, which simultaneously catalyzes the intramolecular rearrangement and reduction of DXP to form 2-C-methyl-d-erythritol 4-phosphate, constitutes a key enzyme of an alternative mevalonate-independent pathway for isopentenyl diphosphate biosynthesis. The dxr gene encoding this enzyme from Escherichia coli was overexpressed as a histidine-tagged protein and characterized in detail. DNA sequencing analysis of the dxr genes from 10 E. coli dxr-deficient mutants revealed base substitution mutations at four points: two nonsense mutations and two amino acid substitutions (Gly(14) to Asp(14) and Glu(231) to Lys(231)). Diethyl pyrocarbonate treatment inactivated DXP reductoisomerase, and subsequent hydroxylamine treatment restored the activity of the diethyl pyrocarbonate-treated enzyme. To characterize these defects, we overexpressed the mutant enzymes G14D, E231K, H153Q, H209Q, and H257Q. All of these mutant enzymes except for G14D were obtained as soluble proteins. Although the purified enzyme E231K had wild-type K(m) values for DXP and NADPH, the mutant enzyme had less than a 0.24% wild-type k(cat) value. K(m) values of H153Q, H209Q, and H257Q for DXP increased to 3.5-, 7.6-, and 19-fold the wild-type value, respectively. These results indicate that Glu(231) of E. coli DXP reductoisomerase plays an important role(s) in the conversion of DXP to 2-C-methyl-d-erythritol 4-phosphate, and that His(153), His(209), and His(257), in part, associate with DXP binding in the enzyme molecule.

#### => d his

(FILE 'HOME' ENTERED AT 11:32:03 ON 13 JAN 2004)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, BIOTECHDS, SCISEARCH' ENTERED AT 11:33:02 ON 13 JAN 2004

L1297 S 1-DEOXY-D-XYLULOSE 5 PHOSPHATE REDUCTOISOMERASE L2162 S L1 AND ISOPRENOID L326 S L2 AND (DNA OR NUCLEIC ACID OR RNA) T.4 12 S L3 AND 2-C METHYL-D-ERYTHRITOL 4-PHOSPHATE L5 10 DUP REM L4 (2 DUPLICATES REMOVED) L6 1 S L5 AND 1990-1998/PY L7 20 DUP REM L3 (6 DUPLICATES REMOVED)  $\Gamma8$ 1 S L7 AND 1993-1998/PY L9 69 S 1-DEOXY-D-XYLULOSE 5 PHOSPHATE REDUCTOISOMERASE AND E. COLI

L10 38 DUP REM L9 (31 DUPLICATES REMOVED)

L11 38 FOCUS L10 1-

=> log y

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 120.94 121.36 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -6.93 -6.93

STN INTERNATIONAL LOGOFF AT 11:44:57 ON 13 JAN 2004